

## SEQUENCE LISTING

<110> LEY, Arthur C. GUTERMAN, Sonia K. MARKLAND, William KENT, Rachel B. ROBERTS, Bruce L. LADNER, Robert C. <120> ITI-D1 KUNITZ DOMAIN MUTANTS AS nHE INHIBITORS <130> LEY=1B <140> 10/038,722 <141> 2002-01-08 <150> US 08/849,406 <151> 1999-07-21 <150> PCT/US95/16349 <151> 1995-12-15 <150> US 08/358,160 <151> 1994-12-16 <160> 129 <170> PatentIn version 3.1 <210> 1 <211> 276 DNA <212> <213> Artificial Sequence <220> <223> IIIsp::bpti::mautreIII (initial fragment) <400> gtgaaaaaat tattattege aatteettta gttgtteett tetattetgg egecegteeg 60 gatttctgtc tcgagccacc atacactggg ccctgcaaag cgcgcatcat ccgctatttc 120 tacaatgcta aagcaggcct gtgccagacc tttgtatacg gtggttgccg tgctaagcgt 180 aacaacttta aatcggccga agattgcatg cgtacctgcg gtggcgccgc tgaaactgtt 240 276 gaaagttgtt tagcaaaacc ccatacagaa aattca <210> 2 <211> 92 <212> PRT <213> Artificial Sequence <220> <223> IIIsp::bpti::mautreIII (initial fragment) <400> 2

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10

15

Gly Ala Lys Glu Asp Ser Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys 20 25 30

Met Gly Met Thr Ser Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys 35 40 45

Glu Thr Phe Gln Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val 50 55 60

Thr Glu Lys Glu Cys Leu Gln Thr Cys Arg Thr Val Gly Ala Ala Glu 65 70 75 80

Thr Val Glu Ser Cys Leu Ala Lys Pro His Thr Glu Asn Ser Phe 85 90 95

<210> 5

<211> 58

<212> PRT

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<223> Consensus Kunitz domain

<400> 5

Arg Pro Asp Phe Cys Leu Leu Pro Ala Glu Thr Gly Pro Cys Arg Ala 1 5 10 15

Met Ile Pro Arg Phe Tyr Tyr Asn Ala Lys Ser Gly Lys Cys Glu Pro 20 25 30

Phe Ile Tyr Gly Gly Cys Gly Gly Asn Ala Asn Asn Phe Lys Thr Glu 35 40 45

Glu Glu Cys Arg Arg Thr Cys Gly Gly Ala 50 55

<210> 6

<211> 58

<212> PRT

<213> Bos Taurus

<400> 6

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Lys Ala 1 5 10 15 Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala 40

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 7 <211> 58 <212> PRT

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<220>

<223> Epi-HNE-1

<400> 7

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Ile Ala

Phe Phe Pro Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 25

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 8

<211> 62

<212> PRT

<213> Artificial Sequence

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<223> Epi-HNE-2

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Glu Ala Glu Ala Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly

Pro Cys Ile Ala Phe Phe Pro Arg Tyr Phe Tyr Asn Ala Lys Ala Gly 20

Leu Cys Gln Thr Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn

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Phe Lys Ser Ala Glu Asp Cys Met Arg Thr Cys Gly Gly Ala
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<210> 9

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<223> EpiNE7

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Met Phe Pro Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 10

<211> 58

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<220>

<223> EpiNE3

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Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Val Gly

Phe Phe Ser Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 25

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

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<211> 58
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Phe Phe Gln Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr
Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala
Glu Asp Cys Met Arg Thr Cys Gly Gly Ala
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<223> EpiNE4

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Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Val Ala

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Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala 55

<210> 13 <211> 58 <212> PRT

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Phe Phe Lys Arg Ser Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 14 <211> 58 <212> PRT

<213> Artificial Sequence

<220>

<223> EpiNE5

<400> 14

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Ile Ala

Phe Phe Gln Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 25

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 15

<211> 58

<213> Artificial Sequence

<220>

<223> EpiNE2

<400> 15

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Ile Ala 10

Leu Phe Lys Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Lys Ser Ala 40

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 16 <211> 58 <212> PRT

<213> Homo sapiens

<400> 16

Lys Glu Asp Ser Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Met Gly

Met Thr Ser Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr 20

Phe Gln Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 40

Lys Asp Cys Leu Gln Thr Cys Arg Thr Val 50

<210> 17

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> BITI-E7-141

<400> 17

Arg Pro Asp Phe Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Val Ala

Met Phe Pro Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Gln Thr 30

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 35 40

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala 50 55

<210> 18

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> MUTT26A

<400> 18

Arg Pro Asp Phe Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Val Ala 1 5 10 15

Met Phe Pro Arg Tyr Phe Tyr Asn Gly Ala Ser Met Ala Cys Gln Thr 20 25 30

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 35 40 45

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala 50 55

<210> 19

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> MUTQE

<400> 19

Arg Pro Asp Phe Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Val Ala 1 5 10 15

Met Phe Pro Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr 20 25 30

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 35 40 45

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala 50

<210> 20

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<211> 58
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<212> PRT

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<220>

<223> MUT1619

<400> 20

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Met Phe Ser Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Gln Thr

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala

<210> 21 <211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> ITI-D1E7

<400> 21

Lys Glu Asp Ser Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Val Ala

Met Phe Pro Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr

Phe Gln Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 35 40

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala

<210> 22

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> AMINO1

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<400> 22
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Lys Glu Asp Phe Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Val Ala

Met Phe Pro Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr

Phe Gln Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala

<210> 23

<211> 58

<212> PRT <213> Artificial Sequence

<220>

<223> AMINO2

<400> 23

Lys Pro Asp Ser Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Val Ala

Met Phe Pro Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr

Phe Gln Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 40

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala

<210> 24

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> MUTP1

<400> 24

Arg Pro Asp Phe Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Ile Gly 1 5

Met Phe Ser Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr 20 25 30

Phe Gln Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 35 40 45

Lys Asp Cys Leu Gln Thr Cys Arg Gly Ala 50 55

<210> 25

<211> 58

<212> PRT

<213> Homo sapiens

<400> 25

Thr Val Ala Ala Cys Asn Leu Pro Ile Val Arg Gly Pro Cys Arg Ala 1 5 10 15

Phe Ile Gln Leu Trp Ala Phe Asp Ala Val Lys Gly Lys Cys Val Leu 20 25 30

Phe Pro Tyr Gly Gly Cys Gln Gly Asn Gly Asn Lys Phe Tyr Ser Glu 35 40 45

Lys Glu Cys Arg Glu Tyr Cys Gly Val Pro 50 55

<210> 26

<211> 56

<212> PRT

<213> Artificial Sequence

<220>

<223> Epi-HNE-3

<400> 26

Ala Ala Cys Asn Leu Pro Ile Val Arg Gly Pro Cys Ile Ala Phe Phe 1 5 10 15

Pro Arg Trp Ala Phe Asp Ala Val Lys Gly Lys Cys Val Leu Phe Pro 20 25 30

Tyr Gly Gly Cys Gln Gly Asn Gly Asn Lys Phe Tyr Ser Glu Lys Glu 35 40 45 Cys Arg Glu Tyr Cys Gly Val Pro

<210> 27

<211> 56

<212> PRT

<213> Artificial Sequence

<220>

<223> Epi-HNE-4

<400> 27

Glu Ala Cys Asn Leu Pro Ile Val Arg Gly Pro Cys Ile Ala Phe Phe

Pro Arg Trp Ala Phe Asp Ala Val Lys Gly Lys Cys Val Leu Phe Pro

Tyr Gly Gly Cys Gln Gly Asn Gly Asn Lys Phe Tyr Ser Glu Lys Glu

Cys Arg Glu Tyr Cys Gly Val Pro

<210> 28 <211> 58 <212> PRT

<213> Homo sapiens

<400> 28

Val Arg Glu Val Cys Ser Glu Gln Ala Glu Thr Gly Pro Cys Arg Ala 5

Met Ile Ser Arg Trp Tyr Phe Asp Val Thr Glu Gly Lys Cys Ala Pro 20

Phe Phe Tyr Gly Gly Cys Gly Gly Asn Arg Asn Asn Phe Asp Thr Glu 40 35

Glu Tyr Cys Met Ala Val Cys Gly Ser Ala 50

<210> 29

<211> 58

<212> PRT

<213> Artificial Sequence

<220> <211> <212> <220>

<223> DPI.1.1 <400> 29

Val Arg Glu Val Cys Ser Glu Gln Ala Tyr Thr Gly Pro Cys Ile Ala

Phe Phe Pro Arg Tyr Tyr Phe Asp Val Thr Glu Gly Lys Cys Gln Thr 25

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Asp Thr Glu

Glu Tyr Cys Met Ala Val Cys Gly Ser Ala

<210> 30 58

PRT

<213> Artificial Sequence

<223> DPI.1.2

<400> 30

Val Arg Glu Val Cys Ser Glu Gln Ala Glu Thr Gly Pro Cys Ile Ala

Met Phe Ser Arg Trp Tyr Phe Asp Val Thr Glu Gly Lys Cys Ala Pro 20

Phe Val Tyr Gly Gly Cys Gly Gly Asn Arg Asn Asn Phe Asp Thr Glu 35

Glu Tyr Cys Met Ala Val Cys Gly Ser Ala 50

<210> 31

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.1.3

<400> 31

Val Arg Glu Val Cys Ser Glu Gln Ala Glu Thr Gly Pro Cys Ile Ala

1 5 10 15

Phe Phe Ser Arg Trp Tyr Phe Asp Val Thr Glu Gly Lys Cys Ala Thr 20 25 30

Phe Val Tyr Gly Gly Cys Met Gly Asn Arg Asn Asn Phe Asp Thr Glu 35 40 45

Glu Tyr Cys Met Ala Val Cys Gly Ser Ala 50 55

<210> 32

<211> 58

<212> PRT

<213> Homo sapiens

<400> 32

Asn Ala Glu Ile Cys Leu Leu Pro Leu Asp Tyr Gly Pro Cys Arg Ala  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Leu Leu Arg Tyr Tyr Tyr Asp Arg Tyr Thr Gln Ser Cys Arg Gln 20 25 30

Phe Leu Tyr Gly Gly Cys Glu Gly Asn Ala Asn Asn Phe Tyr Thr Trp  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Glu Ala Cys Asp Asp Ala Cys Trp Arg Ile 50 55

<210> 33

<211> 58

<212> PRT

<213> Artificial Sequence

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<223> DPI.2.1

<400> 33

Asn Ala Glu Ile Cys Leu Leu Pro Leu Tyr Thr Gly Pro Cys Ile Ala 1 5 10 15

Phe Phe Pro Arg Tyr Tyr Tyr Asp Arg Tyr Thr Gln Ser Cys Gln Thr 20 25 30

Phe Val Tyr Gly Gly Cys Met Gly Asn Ala Asn Asn Phe Tyr Thr Trp 35 40 45

Glu Ala Cys Asp Asp Ala Cys Trp Arg Ile 55

<210> 34

<211> 58

<212> PRT <213> Artificial Sequence

<220>

<223> DPI.2.2

<400> 34

Asn Ala Glu Ile Cys Leu Leu Pro Leu Asp Tyr Gly Pro Cys Ile Ala

Leu Phe Leu Arg Tyr Tyr Tyr Asp Arg Tyr Thr Gln Ser Cys Arg Gln 25

Phe Val Tyr Gly Gly Cys Glu Gly Asn Ala Asn Asn Phe Tyr Thr Trp 40

Glu Ala Cys Asp Asp Ala Cys Trp Arg Ile

<210> 35

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.2.3

<400> 35

Asn Ala Glu Ile Cys Leu Leu Pro Leu Asp Thr Gly Pro Cys Ile Ala

Phe Phe Leu Arg Tyr Tyr Asp Arg Tyr Thr Gln Ser Cys Gln Thr 25

Phe Val Tyr Gly Gly Cys Met Gly Asn Ala Asn Asn Phe Tyr Thr Trp

Glu Ala Cys Asp Asp Ala Cys Trp Arg Ile

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<210> 36
<211> 61
<212> PRT
<213> Homo sapiens
<400> 36
Val Pro Lys Val Cys
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Val Pro Lys Val Cys Arg Leu Gln Val Ser Val Asp Asp Gln Cys Glu  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Gly Ser Thr Glu Lys Tyr Phe Phe Asn Leu Ser Ser Met Thr Cys Glu 20 25 30

Lys Phe Phe Ser Gly Gly Cys His Arg Asn Arg Ile Glu Asn Arg Phe  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Pro Asp Glu Ala Thr Cys Met Gly Phe Cys Ala Pro Lys 50 55 60

<210> 37 <211> 58 <212> PRT <213> Artificial Sequence <220> <223> DPI.3.1

<400> 37

Val Pro Lys Val Cys Arg Leu Gln Val Val Arg Gly Pro Cys Ile Ala 1 5 10 15

Phe Phe Pro Arg Trp Phe Phe Asn Leu Ser Ser Met Thr Cys Val Leu 20 25 30

Phe Pro Tyr Gly Gly Cys Gln Gly Asn Gly Asn Arg Phe Pro Asp Glu 35 40 45

Ala Thr Cys Met Gly Phe Cys Ala Pro Lys 50 55

<210> 38 <211> 61 <212> PRT <213> Artificial Sequence <220> <223> DPI.3.2 <400> 38 Val Pro Lys Val Cys Arg Leu Gln Val Ser Val Asp Asp Gln Cys Ile 10

Gly Ser Phe Glu Lys Tyr Phe Phe Asn Leu Ala Ser Met Thr Cys Glu 25 20

Thr Phe Val Ser Gly Gly Cys His Arg Asn Arg Ile Glu Asn Arg Phe 40

Pro Asp Glu Ala Thr Cys Met Gly Phe Cys Ala Pro Lys

<210> 39

<211> 58 <212> PRT

<213> Artificial Sequence

<220>

<223> DPI.3.3

<400> 39

Val Pro Lys Val Cys Arg Leu Gln Val Val Ala Gly Pro Cys Ile Gly

Phe Phe Lys Arg Tyr Phe Phe Ala Leu Ser Ser Met Thr Cys Glu Thr 20 25

Phe Val Ser Gly Gly Cys His Arg Asn Arg Asn Arg Phe Pro Asp Glu

Ala Thr Cys Met Gly Phe Cys Ala Pro Lys 50 55

<210> 40

<211> 58

<212> PRT

<213> Homo sapiens

<400> 40

Ile Pro Ser Phe Cys Tyr Ser Pro Lys Asp Glu Gly Leu Cys Ser Ala

Asn Val Thr Arg Tyr Tyr Phe Asn Pro Arg Tyr Arg Thr Cys Asp Ala 20 25

Phe Thr Tyr Thr Gly Cys Gly Gly Asn Asp Asn Asn Phe Val Ser Arg

Glu Asp Cys Lys Arg Ala Cys Ala Lys Ala 55

<210> 41

<211> 58 <212> PRT

<213> Artificial Sequence

<220>

<223> DPI.4.1

<400> 41

Ile Pro Ser Phe Cys Tyr Ser Pro Lys Ser Ala Gly Pro Cys Val Ala

Met Phe Pro Arg Tyr Tyr Phe Asn Pro Arg Tyr Arg Thr Cys Glu Thr 25

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Ser Arg

Glu Asp Cys Lys Arg Ala Cys Ala Lys Ala

<210> 42

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.4.2

<400> 42

Ile Pro Ser Phe Cys Tyr Ser Pro Lys Asp Glu Gly Leu Cys Ile Ala

Phe Phe Thr Arg Tyr Tyr Phe Asn Pro Arg Tyr Arg Thr Cys Asp Ala 20 25 30

Phe Thr Tyr Thr Gly Cys Gly Gly Asn Asp Asn Asn Phe Val Ser Arg 35 40 45

Glu Asp Cys Lys Arg Ala Cys Ala Lys Ala 50 55

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<210> 43
<211> 58
<212> PRT
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<223> DPI.4.3
<400> 43
Ile Pro Ser Phe Cys Tyr Ser Pro Lys Asp Thr Gly Pro Cys Ile Ala
Phe Phe Thr Arg Tyr Tyr Phe Asn Pro Arg Tyr Arg Thr Cys Asp Thr
                                25
Phe Val Tyr Gly Gly Cys Gly Gly Asn Asp Asn Asn Phe Val Ser Arg
                            40
Glu Asp Cys Lys Arg Ala Cys Ala Lys Ala
<210> 44
<211> 58
<212> PRT
<213> Homo sapiens
<400> 44
Met His Ser Phe Cys Ala Phe Lys Ala Asp Asp Gly Pro Cys Lys Ala
Ile Met Lys Arg Phe Phe Phe Asn Ile Phe Thr Arg Gln Cys Glu Glu
Phe Ile Tyr Gly Gly Cys Glu Gly Asn Gln Asn Arg Phe Glu Ser Leu
Glu Glu Cys Lys Lys Met Cys Thr Arg Asp
<210> 45
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<223> DPI.5.1

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<400> 45
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Met His Ser Phe Cys Ala Phe Lys Ala Ser Ala Gly Pro Cys Val Ala 1 5 10 15

Met Phe Pro Arg Tyr Phe Phe Asn Ile Phe Thr Arg Gln Cys Glu Thr 20 25 30

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Arg Phe Glu Ser Leu 35 40 45

Glu Glu Cys Lys Lys Met Cys Thr Arg Asp 50 55

<210> 46

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.5.2

<400> 46

Ile Phe Lys Arg Phe Phe Phe Asn Ile Phe Thr Arg Gln Cys Glu Glu 20 25 30

Phe Ile Tyr Gly Gly Cys Glu Gly Asn Gln Asn Arg Phe Glu Ser Leu  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Glu Glu Cys Lys Lys Met Cys Thr Arg Asp

<210> 47

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.5.3

<400> 47

Met His Ser Phe Cys Ala Phe Lys Ala Tyr Thr Gly Pro Cys Ile Ala 1 5 10 15 Phe Phe Lys Arg Phe Phe Phe Asn Ile Phe Thr Arg Gln Cys Glu Thr 25

Phe Ile Tyr Gly Gly Cys Glu Gly Asn Gln Asn Arg Phe Glu Ser Leu 40

Glu Glu Cys Lys Lys Met Cys Thr Arg Asp 55

<210> 48

<211> 58 <212> PRT

<213> Homo sapiens

<400> 48

Lys Pro Asp Phe Cys Phe Leu Glu Glu Asp Pro Gly Ile Cys Arg Gly

Tyr Ile Thr Arg Tyr Phe Tyr Asn Asn Gln Thr Lys Gln Cys Glu Arg

Phe Lys Tyr Gly Gly Cys Leu Gly Asn Met Asn Asn Phe Glu Thr Leu

Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly 55

<210> 49

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.6.1

<400> 49

Lys Pro Asp Phe Cys Phe Leu Glu Glu Ser Ala Gly Pro Cys Val Ala

Met Phe Pro Arg Tyr Phe Tyr Asn Asn Gln Thr Lys Gln Cys Glu Thr 25 20

Phe Val Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Glu Thr Leu 35 40

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Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly
<210> 50
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<213> Artificial Sequence
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<223> DPI.6.2
<400> 50
Lys Pro Asp Phe Cys Phe Leu Glu Glu Asp Pro Gly Ile Cys Val Gly
Tyr Phe Thr Arg Tyr Phe Tyr Asn Asn Gln Thr Lys Gln Cys Glu Arg
Phe Lys Tyr Gly Gly Cys Leu Gly Asn Met Asn Asn Phe Glu Thr Leu
Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly
<210> 51
<211> 58
<212> PRT
<213> Artificial Sequence
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<223> DPI.6.3
<400> 51
Lys Pro Asp Phe Cys Phe Leu Glu Glu Asp Pro Gly Ile Cys Val Gly
Phe Phe Thr Arg Tyr Phe Tyr Asn Asn Gln Thr Lys Gln Cys Glu Arg
         20
                                25
Phe Val Tyr Gly Gly Cys Leu Gly Asn Met Asn Asn Phe Glu Thr Leu
                            40
Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly
                        55
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<210> 52 <211> 58

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<212> PRT
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<213> Artificial Sequence

<220>

<223> DPI.6.4

<400> 52

Lys Pro Asp Phe Cys Phe Leu Glu Glu Asp Pro Gly Ile Cys Val Gly

Phe Phe Thr Arg Tyr Phe Tyr Asn Ala Gln Thr Lys Gln Cys Glu Arg

Phe Val Tyr Gly Gly Cys Leu Gly Asn Met Asn Asn Phe Glu Thr Leu

Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly

<210> 53 <211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.6.5

<400> 53

Lys Pro Asp Phe Cys Phe Leu Glu Glu Asp Pro Gly Pro Cys Val Gly

Phe Phe Gln Arg Tyr Phe Tyr Asn Ala Gln Thr Lys Gln Cys Glu Arg 25

Phe Val Tyr Gly Gly Cys Gln Gly Asn Met Asn Asn Phe Glu Thr Leu 40

Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly

<210> 54

<211> 58

<212> PRT

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<223> DPI.6.6

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<400> 54
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Lys Pro Asp Phe Cys Phe Leu Glu Glu Asp Pro Gly Pro Cys Val Gly

Phe Phe Thr Arg Tyr Phe Tyr Asn Asn Gln Thr Lys Gln Cys Glu Arg .

Phe Val Tyr Gly Gly Cys Gln Gly Asn Met Asn Asn Phe Glu Thr Leu

Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly

<210> 55

<211> 58 <212> PRT

<213> Artificial Sequence

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<223> DPI.6.7

<400> 55

Lys Pro Asp Phe Cys Phe Leu Glu Glu Asp Pro Gly Pro Cys Ile Gly

Phe Phe Pro Arg Tyr Phe Tyr Asn Asn Gln Thr Lys Gln Cys Glu Arg 25

Phe Val Tyr Gly Gly Cys Gln Gly Asn Met Asn Asn Phe Glu Thr Leu 40

Glu Glu Cys Lys Asn Ile Cys Glu Asp Gly 55

<210> 56

<211> 58

<212> PRT

<213> Homo sapiens

Gly Pro Ser Trp Cys Leu Thr Pro Ala Asp Arg Gly Leu Cys Arg Ala

Asn Glu Asn Arg Phe Tyr Tyr Asn Ser Val Ile Gly Lys Cys Arg Pro

Phe Lys Tyr Ser Gly Cys Gly Gly Asn Glu Asn Asn Phe Thr Ser Lys 40

Gln Glu Cys Leu Arg Ala Cys Lys Lys Gly . 55

<210> 57

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<212> PRT <213> Artificial Sequence

<220>

<223> DPI.7.1

<400> 57

Gly Pro Ser Trp Cys Leu Thr Pro Ala Val Arg Gly Pro Cys Ile Ala

Phe Phe Pro Arg Trp Tyr Tyr Asn Ser Val Ile Gly Lys Cys Val Leu

Phe Pro Tyr Gly Gly Cys Gln Gly Asn Gly Asn Asn Phe Thr Ser Lys

Gln Glu Cys Leu Arg Ala Cys Lys Gly

<210> 58

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.7.2

<400> 58

Gly Pro Ser Trp Cys Leu Thr Pro Ala Asp Arg Gly Leu Cys Val Ala 5

Asn Phe Asn Arg Phe Tyr Tyr Asn Ser Val Ile Gly Lys Cys Arg Pro

Phe Lys Tyr Ser Gly Cys Gly Gly Asn Glu Asn Asn Phe Thr Ser Lys

Gln Glu Cys Leu Arg Ala Cys Lys Lys Gly

50 55

<210> 59

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.7.3

<400> 59

Gly Pro Ser Trp Cys Leu Thr Pro Ala Asp Arg Gly Leu Cys Val Ala

Phe Phe Asn Arg Phe Tyr Tyr Asn Ser Val Ile Gly Lys Cys Arg Pro

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Gln Glu Cys Leu Arg Ala Cys Lys Lys Gly

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<211> 58 <212> PRT

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<223> DPI.7.4

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<223> DPI.7.5

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Phe Val Tyr Gly Gly Cys Gly Gly Asn Glu Asn Asn Phe Ala Ser Lys 35 40 45

Gln Glu Cys Leu Arg Ala Cys Lys Lys Gly
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<210> 62

<211> 58

<212> PRT

<213> Homo sapiens

<400> 62

Glu Thr Asp Ile Cys Lys Leu Pro Lys Asp Glu Gly Thr Cys Arg Asp 1  $\phantom{-}5\phantom{+}10\phantom{+}15\phantom{+}15$ 

Phe Ile Leu Lys Trp Tyr Tyr Asp Pro Asn Thr Lys Ser Cys Ala Arg 20 25 30

Phe Trp Tyr Gly Gly Cys Gly Gly Asn Glu Asn Lys Phe Gly Ser Gln 35 40 45

Lys Glu Cys Glu Lys Val Cys Ala Pro Val

<210> 63

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.8.1

<400> 63

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Phe Pro Tyr Gly Gly Cys Gln Gly Asn Gly Asn Lys Phe Gly Ser Gln 35 40 45

Lys Glu Cys Glu Lys Val Cys Ala Pro Val 50 55

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<211> 58

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<211> 58

<212> PRT

<213> Artificial Sequence

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<400> 65

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Lys Glu Cys Glu Lys Val Cys Ala Pro Val 50 55

<210> 66

<211> 58

<212> PRT

<213> Homo sapiens

<400> 66

Leu Pro Asn Val Cys Ala Phe Pro Met Glu Lys Gly Pro Cys Gln Thr
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Tyr Met Thr Arg Trp Phe Phe Asn Phe Glu Thr Gly Glu Cys Glu Leu 20 25 30

Phe Ala Tyr Gly Gly Cys Gly Gly Asn Ser Asn Asn Phe Leu Arg Lys 35 40 45

Glu Lys Cys Glu Lys Phe Cys Lys Phe Thr 50 55

<210> 67

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> DPI.9.1

<400> 67

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Glu Lys Cys Glu Lys Phe Cys Lys Phe Thr 50 55

<210> 68

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<sup>&</sup>lt;400> 71

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<sup>&</sup>lt;211> 8584

<sup>&</sup>lt;212> DNA

<sup>&</sup>lt;213> Artificial Sequence

<sup>&</sup>lt;220>

<sup>&</sup>lt;223> Plasmid pHIL-D2 (MFalphaPrePro::EPI-HNE-3) (Table 251)

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ggcgtgctgc tagcgctata tgcgttgatg caatttctat gcgcacccgt tctcggagca 1980 ctgtccgacc gctttggccg ccgcccagtc ctgctcgctt cgctacttgg agccactatc 2040 gactacgcga tcatggcgac cacacccgtc ctgtggatct atcgaatcta aatgtaagtt 2100 aaaatotota aataattaaa taagtoocag tttotooata ogaacottaa cagcattgog 2160 gtgagcatct agaccttcaa cagcagccag atccatcact gcttggccaa tatgtttcag 2220 teceteagga gttaegtett gtgaagtgat gaaettetgg aaggttgeag tgttaaetee 2280 gctgtattga cgggcatatc cgtacgttgg caaagtgtgg ttggtaccgg aggagtaatc 2340 tocacaacto totggagagt aggcaccaac aaacacagat ccagcgtgtt gtacttgate 2400 aacataagaa gaagcattct cgatttgcag gatcaagtgt tcaggagcgt actgattgga 2460 catttccaaa gcctgctcgt aggttgcaac cgatagggtt gtagagtgtg caatacactt 2520 gogtacaatt toaaccottg goaactgoac agettggttg tgaacagoat ottoaattot 2580 ggcaagctcc ttgtctgtca tatcgacagc caacagaatc acctgggaat caataccatg 2640 ttcagcttga gcagaaggtc tgaggcaacg aaatctggat cagcgtattt atcagcaata 2700 'actagaactt cagaaggccc agcaggcatg tcaatactac acagggctga tgtgtcattt 2760 tgaaccatca tettggcage agtaacgaae tggttteetg gaccaaatat tttgtcacae 2820 ttaggaacag tttctgttcc gtaagccata gcagctactg cctgggcgcc tcctgctagc 2880 acgatacact tagcaccaac cttgtgggca acgtagatga cttctggggt aagggtacca 2940 teettettag gtggagatge aaaaacaatt tetttgcaae cagcaaettt ggcaggaaca 3000 cccagcatca gggaagtgga aggcagaatt gcggttccac caggaatata gaggccaact 3060 tteteaatag gtettgeaa'a acgagageag actacaceag ggeaagtete aacttgeaac 3120 gtctccgtta gttgagcttc atggaatttc ctgacgttat ctatagagag atcaatggct 3180 ctcttaacgt tatctggcaa ttgcataagt tcctctggga aaggagcttc taacacaggt 3240 gtcttcaaag cgactccatc aaacttggca gttagttcta aaagggcttt gtcaccattt 3300 tgacgaacat tgtcgacaat tggtttgact aattccataa tctgttccgt tttctggata 3360 ggacgacgaa gggcatcttc aatttcttgt gaggaggcct tagaaacgtc aattttgcac 3420 aattcaatac gaccttcaga agggacttct ttaggtttgg attcttcttt aggttgttcc 3480 ttggtgtatc ctggcttggc atctcctttc cttctagtga cctttaggga cttcatatcc 3540 aggtttctct ccacctcgtc caacgtcaca ccgtacttgg cacatctaac taatgcaaaa 3600 taaaataagt cagcacattc ccaggctata tcttccttgg atttagcttc tgcaagttca 3660

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i ?

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                                                                       7260
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                                                                      7500
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                                                                      7920
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                                                                     8280
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                                                                     8520
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                                                                     8580
gcag
                                                                     8584
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<sup>&</sup>lt;210> 72

<sup>&</sup>lt;211> 141

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Artificial Sequence

<sup>&</sup>lt;220>

<sup>&</sup>lt;223> Plasmid pHIL-D2 (MFalphaPrePro::EPI-HNE-3) (Table 251)

<400> 72

Met Arg Phe Pro Ser Ile Phe Thr Ala Val Leu Phe Ala Ala Ser Ser 10 10 15

Ala Leu Ala Ala Pro Val Asn Thr Thr Glu Asp Glu Thr Ala Gln 20 25 30

Ile Pro Ala Glu Ala Val Ile Gly Tyr Ser Asp Leu Glu Gly Asp Phe 35 40 45

Asp Val Ala Val Leu Pro Phe Ser Asn Ser Thr Asn Asn Gly Leu Leu 50 55 60

Phe Ile Asn Thr Thr Ile Ala Ser Ile Ala Ala Lys Glu Glu Gly Val 70 75 80

Ser Leu Asp Lys Arg Ala Ala Cys Asn Leu Pro Ile Val Arg Gly Pro 85 90 95

Cys Ile Ala Phe Phe Pro Arg Trp Ala Phe Asp Ala Val Lys Gly Lys 100 105 110

Tyr Ser Glu Lys Glu Cys Arg Glu Tyr Cys Gly Val Pro 130 135 140

<210> 73

<211> 444

<212> DNA

<213> Artificial Sequence

<220>

<223> BstBI-AatII-EcoRI cassette for expression of Epi-HNE-4 (Table 252)

<400> 73

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ttggctgctc cagttaacac cactactgaa gacgagactg ctcaaattcc tgctgaggct 120
gtcatcggtt actctgactt ggaaggtgac ttcgacgtcg ctgtttgcc attctctaac 180
tctactaaca acggtttgtt gttcatcaac actaccatcg cttctatcgc tgctaaggag 240
gaaggtgttt ccttggacaa gagagaggct tgtaacttgc caatcgtcag aggtccatgc 300

attgctttct tcccaagatg ggctttcgac gctgttaagg gtaagtgcgt cttgttccca 360 tacggtggtt gtcaaggtaa cggtaacaag ttctactctg agaaggagtg tagagagtac 420 444 tgtggtgttc catagtaaga attc <210> 74 <211> 141 <212> PRT <213> Artificial Sequence <220> BstBI-AatII-EcoRI cassette for expression of Epi-HNE-4 (Table 252 <400> 74 . Met Arg Phe Pro Ser Ile Phe Thr Ala Val Leu Phe Ala Ala Ser Ser Ala Leu Ala Ala Pro Val Asn Thr Thr Glu Asp Glu Thr Ala Gln 20 Ile Pro Ala Glu Ala Val Ile Gly Tyr Ser Asp Leu Glu Gly Asp Phe. 35 40 Asp Val Ala Val Leu Pro Phe Ser Asn Ser Thr Asn Asn Gly Leu Leu 50 55 Phe Ile Asn Thr Thr Ile Ala Ser Ile Ala Ala Lys Glu Glu Gly Val 65 70 Ser Leu Asp Lys Arg Glu Ala Cys Asn Leu Pro Ile Val Arg Gly Pro 85 Cys Ile Ala Phe Phe Pro Arg Trp Ala Phe Asp Ala Val Lys Gly Lys 100 105 Cys Val Leu Phe Pro Tyr Gly Gly Cys Gln Gly Asn Gly Asn Lys Phe 115 Tyr Ser Glu Lys Glu Cys Arg Glu Tyr Cys Gly Val Pro <210> 75 <211> 8590 <212> DNA

<213> Artificial Sequence

<220>

<223> pD2pick (MFalphaPrePro::EPI-NHE-3) circular dsDNA (Table 253) <400> 75 agatcgcggc cgcgatctaa catccaaaga cgaaaggttg aatgaaacct tttttgccatc 60 120 cgacatccac aggtccattc tcacacataa gtgccaaacg caacaggagg ggatacacta gcagcagacc gttgcaaacg caggacctcc actcctcttc tcctcaacac ccacttttgc 180 categaaaaa ccageeeagt tattgggett gattggaget egeteattee aatteettet 240 attaggctac taacaccatg actttattag cctgtctatc ctggcccccc tggcgaggtc 300 360 atgtttgttt atttccgaat gcaacaagct ccgcattaca cccgaacatc actccagatg agggetttet gagtgtgggg teaaatagtt teatgtteee aaatggeeea aaactgaeag 420 tttaaacgct gtcttggaac ctaatatgac aaaagcgtga tctcatccaa gatgaactaa 480 gtttggttcg ttgaaatgct aacggccagt tggtcaaaaa gaaacttcca aaagtcgcca 540 600 taccgtttgt cttgtttggt attgattgac gaatgctcaa aaataatctc attaatgctt 660 agegeagtet etetateget tetgaaceeg gtggeacetg tgeegaaaeg caaatgggga 720 aacaacccgc tttttggatg attatgcatt gtcctccaca ttgtatgctt ccaagattct ggtgggaata ctgctgatag cctaacgttc atgatcaaaa tttaactgtt ctaaccccta 780 840 % cttgacaggc aatatataaa cagaaggaag ctgccctgtc ttaaaccttt ttttttatca 900 tcattattag cttactttca taattgcgac tggttccaat tgacaagctt ttgattttaa 960 cgacttttaa cgacaacttg agaagatcaa aaaacaacta attattcgaa acgatgagat 1020 teccatetat etteaetget gttttgtteg etgetteete tgetttgget geteeagtta acaccactac tgaagacgag actgctcaaa ttcctgctga ggctgtcatc ggttactctg 1080 acttggaagg tgacttcgac gtcgctgttt tgccattctc taactctact aacaacggtt 1140 1200 tgttgttcat caacactacc atcgcttcta tcgctgctaa ggaggaaggt gtttccttgg acaagagage tgcttgtaac ttgccaatcg tcagaggtcc atgcattgct ttcttcccaa 1260 gatgggettt egaegetgtt aagggtaagt gegtettgtt eeeataeggt ggttgteaag 1320 gtaacggtaa caagttctac tctgagaagg agtgtagaga gtactgtggt gttccatagt 1380 aagaattcgc cttagacatg actgttcctc agttcaagtt gggcattacg agaagaccgg 1440 tettgetaga ttetaateaa gaggatgtea gaatgeeatt tgeetgagag atgeaggett 1500

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<220>

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<223> Xaa is Asn or Gly
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<221> misc_feature
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<223> Xaa is any amino acid
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<400> 86
Cys Xaa Xaa Xaa Xaa Xaa Gly Xaa Cys Xaa Xaa Xaa Xaa Xaa Xaa
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Cys Xaa Xaa Phe Xaa Xaa
            20
                                 25
Gly Cys Xaa Xaa Xaa Xaa Xaa Aaa Phe Xaa Xaa Xaa Xaa Cys Xaa
                            40
        35
Xaa Xaa Cys
    50
<210> 87
<211> 58
<212> PRT
<213> Bos Taurus
<400> 87
Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Lys Ala
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Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 88

<211> 58 <212> PRT

<213> Artificial Sequence

<220>

<223> Engineered B-PTI from MARK87

<400> 88

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Thr Lys Ala

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Thr Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 89

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> Engineered B-PTI from MARK87

<400> 89

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Ala Lys Ala

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 20 25

Phe Val Tyr Gly Gly Ala Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala 35 40

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala
50 55

<210> 90

<211> 67

<212> PRT

<213> Bos taurus

<400> 90

Phe Gln Thr Pro Pro Asp Leu Cys Gln Leu Pro Gln Ala Arg Gly Pro 1 5 10 15

Cys Lys Ala Ala Leu Leu Arg Tyr Phe Tyr Asn Ser Thr Ser Asn Ala 20 25 30

Cys Glu Pro Phe Thr Tyr Gly Gly Cys Gln Gly Asn Asn Asn Phe 35 40 45

Glu Thr Thr Glu Met Cys Leu Arg Ile Cys Glu Pro Pro Gln Gln Thr 50 60

Asp Lys Ser 65

<210> 91

<211> 60

<212> PRT

<213> Bos Taurus

<400> 91

Thr Glu Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys

10 15

Lys Ala Ala Met Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Phe Cys 20 25 30

Glu Thr Phe Val Tyr Gly Gly Cys Arg Ala Lys Ser Asn Asn Phe Lys 35 40 45

Ser Ala Glu Asp Cys Met Arg Thr Cys Gly Gly Ala 50 55 60

<210> 92

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> Semisynthetic BPTI, TSCH87

<400> 92

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Val Ala

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser $\cdot$ Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 93 <211> 58 <212> PRT

<213> Artificial Sequence

<220>

<223> Semisynthetic BPTI, TSCH87

<400> 93

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Gly Ala

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala 50

<210> 94

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> Semisynthetic BPTI, TSCH87

<400> 94

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Ala Ala

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 95 <211> 58 <212> PRT

<213> Artificial Sequence

<220>

<223> Semisynthetic BPTI, TSCH87

<400> 95

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Leu Ala

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 25

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala 40

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 96

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> Semisynthetic BPTI, TSCH87

<400> 96

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Ile Ala 10

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 20 25 30

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala 35 40 45

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala 50 55

<210> 97

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> Engineered BPTI, AUER87

<400> 97

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Lys Ala
1 5 10 15

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 20 25 30

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala 35 40 45

Glu Asp Cys Glu Arg Thr Cys Gly Gly Ala 50 55

<210> 98

<211> 60

<212> PRT

<213> Dendroaspis polylepis polylepis

<400> 98

Gln Pro Leu Arg Lys Leu Cys Ile Leu His Arg Asn Pro Gly Arg Cys
1 10 15

Tyr Gln Lys Ile Pro Ala Phe Tyr Tyr Asn Gln Lys Lys Lys Gln Cys 20 25 30

Glu Gly Phe Thr Trp Ser Gly Cys Gly Gly Asn Ser Asn Arg Phe Lys 35 40 45

Thr Ile Glu Glu Cys Arg Arg Thr Cys Ile Arg Lys

55 50 60

<210> 99

<211> 57

<212> PRT

<213> Dendroaspis polylepis polylepis

<400> 99

Ala Ala Lys Tyr Cys Lys Leu Pro Leu Arg Ile Gly Pro Cys Lys Arg

Lys Ile Pro Ser Phe Tyr Tyr Lys Trp Lys Ala Lys Gln Cys Leu Pro

Phe Asp Tyr Ser Gly Cys Gly Gly Asn Ala Asn Arg Phe Lys Thr Ile

Glu Glu Cys Arg Arg Thr Cys Val Gly

<210> 100 <211> 57

<212> PRT

<213> Hemachatus hemachates

<400> 100

Arg Pro Asp Phe Cys Glu Leu Pro Ala Glu Thr Gly Leu Cys Lys Ala 5

Tyr Ile Arg Ser Phe His Tyr Asn Leu Ala Ala Gln Gln Cys Leu Gln 20

Phe Ile Tyr Gly Gly Cys Gly Gly Asn Ala Asn Arg Phe Lys Thr Ile 40

Asp Glu Cys Arg Arg Thr Cys Val Gly 50

<210> 101

<211> 57

<212> PRT

<213> Naja nivea

<400> 101

Arg Pro Arg Phe Cys Glu Leu Pro Ala Glu Thr Gly Leu Cys Lys Ala 10

Arg Ile Arg Ser Phe His Tyr Asn Arg Ala Ala Gln Gln Cys Leu Glu 20 25 30

Phe Ile Tyr Gly Gly Cys Gly Gly Asn Ala Asn Arg Phe Lys Thr Ile  $35 \hspace{1.5cm} 40 \hspace{1.5cm} 45$ 

Asp Glu Cys His Arg Thr Cys Val Gly 50

<210> 102

<211> 60

<212> PRT

<213> Vipera russelli

<400> 102

His Asp Arg Pro Thr Phe Cys Asn Leu Pro Pro Glu Ser Gly Arg Cys  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Arg Gly His Ile Arg Ile Tyr Tyr Asn Leu Glu Ser Asn Lys Cys 20 25 30

Lys Val Phe Phe Tyr Gly Gly Cys Gly Gly Asn Ala Asn Asn Phe Glu 35 40 45

Thr Arg Asp Glu Cys Arg Glu Thr Cys Gly Gly Lys 50 55 60

<210> 103

<211> 64

<212> PRT

<213> Caretta sp.

<400> 103

Glx Gly Asp Lys Arg Asp Ile Cys Arg Leu Pro Pro Glu Gln Gly Pro 1 5 10 15

Cys Lys Gly Arg Leu Pro Arg Tyr Phe Tyr Asn Pro Ala Ser Arg Met 20 25 30

Cys Glu Ser Phe Ile Tyr Gly Gly Cys Lys Gly Asn Lys Asn Asn Phe 35 40 45

Lys Thr Lys Ala Glu Cys Val Arg Ala Cys Arg Pro Pro Glu Arg Pro 50 55 60

<210> 104

<211> 58

<212> PRT

<213> Helix pomatia

<400> 104

Glx Gly Arg Pro Ser Phe Cys Asn Leu Pro Ala Glu Thr Gly Pro Cys .  $10 ext{1}$ 

Lys Ala Ser Ile Arg Gln Tyr Tyr Tyr Asn Ser Lys Ser Gly Gly Cys 20 25 30

Gln Gln Phe Ile Tyr Gly Gly Cys Arg Gly Asn Gln Asn Arg Phe Asp  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Thr Thr Gln Gln Cys Gln Gly Val Cys Val
50 55

<210> 105

<211.> 57

<212> PRT

<213> Dendroaspis angusticeps

<400> 105

Ala Ala Lys Tyr Cys Lys Leu Pro Val Arg Tyr Gly Pro Cys Lys 1 5 10 15

Lys Phe Pro Ser Phe Tyr Tyr Asn Trp Lys Ala Lys Gln Cys Leu Pro 20 25 30

Phe Asn Tyr Ser Gly Cys Gly Gly Asn Ala Asn Arg Phe Lys Thr Ile  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Glu Glu Cys Arg Arg Thr Cys Val Gly 50

<210> 106

<211> 59

<212> PRT

<213> Dendroaspis angusticeps

<400> 106

Glx Pro Arg Arg Lys Leu Cys Ile Leu His Arg Asn Pro Gly Arg Cys
1 10 15

Tyr Asp Lys Ile Pro Ala Phe Tyr Tyr Asn Gln Lys Lys Lys Gln Cys 20 25 30

Glu Arg Phe Asp Trp Ser Gly Cys Gly Gly Asn Ser Asn Arg Phe Lys 35 40 45

Thr Ile Glu Glu Cys Arg Arg Thr Cys Ile Gly

<210> 107

<211> 57

<212> PRT

<213> Dendroaspis polylepis polylepis

<400> 107

Arg Pro Tyr Ala Cys Glu Leu Ile Val Ala Ala Gly Pro Cys Met Phe 1  $\phantom{-}5\phantom{+}10\phantom{+}10\phantom{+}15\phantom{+}10$ 

Phe Ile Ser Ala Phe Tyr Tyr Ser Lys Gly Ala Asn Lys Cys Tyr Pro 20 25 30

Phe Thr Tyr Ser Gly Cys Arg Gly Asn Ala Asn Arg Phe Lys Thr Ile 35 40 45

Glu Glu Cys Arg Arg Thr Cys Val Val 50 55

. <210> 108

<211> 59

<212> PRT

<213> Dendroaspis polylepis polylepis

<400> 108

Leu Gln His Arg Thr Phe Cys Lys Leu Pro Ala Glu Pro Gly Pro Cys 1 10 15

Lys Ala Ser Ile Pro Ala Phe Tyr Tyr Asn Trp Ala Ala Lys Lys Cys 20 25 30

Gln Leu Phe His Tyr Gly Gly Cys Lys Gly Asn Ala Asn Arg Phe Ser 35 40 45

Thr Ile Glu Lys Cys Arg His Ala Cys Val Gly 50

<210> 109

<211> 61

<212> PRT

<213> Vipera ammodytes

<400> 109

Glx Asp His Pro Lys Phe Cys Tyr Leu Pro Ala Asp Pro Gly Arg Cys

Lys Ala His Ile Pro Arg Phe Tyr Tyr Asp Ser Ala Ser Asn Lys Cys

Asn Lys Phe Ile Tyr Gly Gly Cys Pro Gly Asn Ala Asn Asn Phe Lys

Thr Trp Asp Glu Cys Arg Gln Thr Cys Gly Ala Ser Ala

<210> 110

<211> 62 <212> PRT

<213> Vipera ammodytes

<400> 110

Arg Asp Arg Pro Lys Phe Cys Tyr Leu Pro Ala Asp Pro Gly Arg Cys

Leu Ala Tyr Met Pro Arg Phe Tyr Tyr Asn Pro Ala Ser Asn Lys Cys 20

Glu Lys Phe Ile Tyr Gly Gly Cys Arg Gly Asn Ala Asn Asn Phe Lys 35

Thr Trp Asp Glu Cys Arg His Thr Cys Val Ala Ser Gly Ile 50

<210> 111

<211> 62

<212> PRT

<213> Bungarus fasciatus

<400> 111

Lys Asn Arg Pro Thr Phe Cys Asn Leu Leu Pro Glu Thr Gly Arg Cys

Asn Ala Leu Ile Pro Ala Phe Tyr Tyr Asn Ser His Leu His Lys Cys 20 25 30

Gln Lys Phe Asn Tyr Gly Gly Cys Gly Gly Asn Ala Asn Asn Phe Lys  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Thr Ile Asp Glu Cys Gln Arg Thr Cys Ala Ala Lys Tyr Gly 50 55 60

<210> 112

<211> 59

<212> PRT

<213> Anemonia sulcata

<400> 112

Ile Asn Gly Asp Cys Glu Leu Pro Lys Val Val Gly Pro Cys Arg Ala 1 5 10 15

Arg Phe Pro Arg Tyr Tyr Tyr Asn Ser Ser Ser Lys Arg Cys Glu Lys 20 25 30

Phe Ile Tyr Gly Gly Cys Gly Gly Asn Ala Asn Asn Phe His Thr Leu 35 40 45

Glu Glu Cys Glu Lys Val Cys Gly Val Arg Ser 50 55

<210> 113

<211> 56

<212> PRT

<213> Homo sapiens

<400> 113

Lys Glu Asp Ser Cys Gln Leu Gly Tyr Ser Ala Gly Pro Cys Met Gly 1  $\phantom{-}$  5  $\phantom{-}$  10  $\phantom{-}$  15

Met Thr Ser Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr 20 25 30

Phe Gln Tyr Gly Gly Cys Met Gly Asn Gly Asn Asn Phe Val Thr Glu 35 40 45

Lys Glu Cys Leu Gln Thr Cys Arg 50 55 <210> 114

<211> 61

<212> PRT

<213> Homo sapiens

<400> 114

Thr Val Ala Ala Cys Asn Leu Pro Val Ile Arg Gly Pro Cys Arg Ala 1 5 10 15

Phe Ile Gln Leu Trp Ala Phe Asp Ala Val Lys Gly Lys Cys Val Leu 20 25 30

Phe Pro Tyr Gly Gly Cys Gln Gly Asn Gly Asn Lys Phe Tyr Ser Glu 35 40 45

Lys Glu Cys Arg Glu Tyr Cys Gly Val Pro Gly Asp Glu 50 55 60

<210> 115

<211> 60

<212> PRT

<213> Bungarus multicinctus

<400> 115

Arg Gln Arg His Arg Asp Cys Asp Lys Pro Pro Asp Lys Gly Asn Cys  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Gly Pro Val Arg Ala Phe Tyr Tyr Asp Thr Arg Leu Lys Thr Cys Lys 20 25 30

Ala Phe Gln Tyr Arg Gly Cys Asp Gly Asp His Gly Asn Phe Lys Thr 35 40 45

Glu Thr Leu Cys Arg Cys Glu Cys Leu Val Tyr Pro 50 55 60

<210> 116

<211> 60

<212> PRT

<213> Bungarus multicinctus

<400> 116

Arg Lys Arg His Pro Asp Cys Asp Lys Pro Pro Asp Thr Lys Ile Cys
1 10 15

Gln Thr Val Arg Ala Phe Tyr Tyr Lys Pro Ser Ala Lys Arg Cys Val

20 25 30

Gln Phe Arg Tyr Gly Gly Cys Asp Gly Asp His Gly Asn Phe Lys Ser 35 40 45

Asp His Leu Cys Arg Cys Glu Cys Glu Leu Tyr Arg 50 55 60

<210> 117

<211> 58

<212> PRT

<213> Bos taurus

<400> 117

Lys Met Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Phe Cys Glu Thr 20 25 30

Phe Val Tyr Gly Gly Cys Lys Ala Lys Ser Asn Asn Phe Arg Ser Ala 35 40 45

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala 50 55

<210> 118

<211> 61

<212> PRT

<213> Tachypleus tridentatus

<400> 118

Thr Glu Arg Gly Phe Leu Asp Cys Thr Ser Pro Pro Val Thr Gly Pro 1 5 10 15

Cys Arg Ala Gly Phe Lys Arg Tyr Asn Tyr Asn Thr Arg Thr Lys Gln
20 25 30

Cys Glu Pro Phe Lys Tyr Gly Gly Cys Lys Gly Asn Gly Asn Arg Tyr 35 40 45

Lys Ser Glu Gln Asp Cys Leu Asp Ala Cys Ser Gly Phe 50 55 60

<210> 119

<211> 63

<212> PRT

<213> Bombyx mori

<400> 119

Asp Glu Pro Thr Thr Asp Leu Pro Ile Cys Glu Gln Ala Phe Gly Asp 1 5 10 15

Ala Gly Leu Cys Phe Gly Tyr Met Lys Leu Tyr Ser Tyr As<br/>n Gl<br/>n Glu 20 25 30

Thr Lys Asn Cys Glu Glu Phe Ile Tyr Gly Gly Cys Gln Gly Asn Asp  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Asn Arg Phe Ser Thr Leu Ala Glu Cys Glu Gln Lys Cys Ile Asn 50 55 60

<210> 120

<211> 56

<212> PRT

<213> Bos taurus

<400> 120

Lys Ala Asp Ser Cys Gln Leu Asp Tyr Ser Gln Gly Pro Cys Leu Gly
1 5 10 15

Leu Phe Lys Arg Tyr Phe Tyr Asn Gly Thr Ser Met Ala Cys Glu Thr 20 25 30

Phe Leu Tyr Gly Gly Cys Met Gly Asn Leu Asn Asn Phe Leu Ser Gln 35 40 45

Lys Glu Cys Leu Gln Thr Cys Arg

<210> 121

<211> 61

<212> PRT

<213> Bos taurus

<400> 121

Thr Val Glu Ala Cys Asn Leu Pro Ile Val Gln Gly Pro Cys Arg Ala
1 5 10 15

Phe Ile Gln Leu Trp Ala Phe Asp Ala Val Lys Gly Lys Cys Val Arg 20 25 30 Phe Ser Tyr Gly Gly Cys Lys Gly Asn Gly Asn Lys Phe Tyr Ser Gln 35 40 45

Lys Glu Cys Lys Glu Tyr Cys Gly Ile Pro Gly Glu Ala 50 55 60

<210> 122

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> Engineered BPTI (KR15, ME52)

<400> 122

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Arg Ala 1 5 10 15

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Thr 20 25 30

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala . 35 40 45

Glu Asp Cys Glu Arg Thr Cys Gly Gly Ala 50 55

<210> 123

<211> 59

<212> PRT

<213> Artificial Sequence

<220>

<223> Isoaprotinin G-1

<400> 123

Ala Arg Met Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln 20 25 30

Pro Phe Val Tyr Gly Gly Cys Arg Ala Lys Ser Asn Asn Phe Lys Ser 35 40 45

Ala Glu Asp Cys Met Arg Thr Cys Gly Gly Ala 55

<210> 124

<211> 58

<212> PRT

<213> Artificial Sequence

<220>

<223> Isoaprotinin 2

<400> 124

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Lys Ala

Arg Ile Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Pro

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ser

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 125 <211> 58 <212> PRT

<213> Artificial Sequence

<220>

<223> Isoaprotinin G-2

<400> 125

Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Lys Ala

Arg Met Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Leu Cys Gln Pro 20 25

Phe Val Tyr Gly Gly Cys Arg Ala Lys Arg Asn Asn Phe Lys Ser Ala 40

Glu Asp Cys Met Arg Thr Cys Gly Gly Ala

<210> 126

<211> 58

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<212> PRT
<213> Artificial Sequence
<220>
<223> Isoaprotinin 1
<400> 126
Arg Pro Asp Phe Cys Leu Glu Pro Pro Tyr Thr Gly Pro Cys Lys Ala
Lys Met Ile Arg Tyr Phe Tyr Asn Ala Lys Ala Gly Phe Cys Glu Thr
Phe Val Tyr Gly Gly Cys Lys Ala Lys Ser Asn Asn Phe Arg Ser Ala
Glu Asp Cys Met Arg Thr Cys Gly Gly Ala
<210> 127
<211> 11
<212> DNA
<213> Artificial Sequence
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<223> PfMI restriction site
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<221> misc_feature
<222> (4)..(8)
<223> n is a, c, g or t
<400> 127
                                                                     11
ccannnnntg g
<210> 128
<211> 15
<212> DNA
<213> Artificial Sequence
<220>
<223> XcmI restriction site
<220>
<221> misc_feature
<222> (4)..(12)
<223> n is a, c, g or t
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<400> 128

ccannnnnn nntgg

15

<210> 129
<211> 9
<212> PRT
<213> Artificial Sequence
<220>
<223> amino acids 13-21 of EpiNE alpha
<400> 129

Pro Cys Val Ala Met Phe Gln Arg Tyr